10/918,318 (amended)

L20 ANSWER 121 OF 181 CAPLUS COPYRIGHT 2008 ACS on STN

AN 1986:68665 CAPLUS

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OREF 104:10985a,10988a

TI (o- and p-Nitrobenzyloxycarbonyl)-5-fluorouracil derivatives as potential conjugated bioreductive alkylating agents

AU Lin, Tai Shun; Wang, Lin; Antonini, Ippolito; Cosby, Lucille A.; Shiba, David A.; Kirkpatrick, D. Lynn; Sartorelli, Alan C.

CS Sch. Med., Yale Univ., New Haven, CT, 06510, USA

SO Journal of Medicinal Chemistry (1986), 29(1), 84-9 CODEN: JMCMAR; ISSN: 0022-2623

DT Journal

LA English

OS CASREACT 104:68665

AB A series of 5-fluorouracil derivs. I (R, R1 = H, CO2CH2C6H4NO2-4, CO2CH2C6H4NO2-2) were synthesized by reacting o- or p-O2NC6H4CH2O2CCl with 5-fluorouracil in the presence of Et3N in DMF or Me2SO. The reductive activation of these agents was hypothesized to generate a reactive methide and 5-fluorouracil, two components that are capable to synergistic interaction through complementary inhibition. Measurement of the surviving fractions of EMT6 tumor cells treated with these agents in culture under conditions of hypoxia and aerobiosis resulted in equal cell kill regardless of the state of oxygenation. I (R = H, R1 = CO2CH2C6H4NO2-4) appeared to be superior to 5-fluorouracil in prolonging the survival time of mice bearing i.p. implants of the P388 leukemia and Sarcoma 180.

RN 98653-09-9 CAPLUS

CN 2,4(1H,3H)-Pyrimidinedione, 5-fluoro-3-[(2-nitrophenyl)methyl]- (CA INDEX NAME)

$$\begin{array}{c|c}
H & O \\
N & CH_2 \\
O & O_2N
\end{array}$$

RN 98653-10-2 CAPLUS

CN 2,4(1H,3H)-Pyrimidinedione, 5-fluoro-3-[(3-nitrophenyl)methyl]- (CA INDEX NAME)

10/918,318 (amended)

RN 98653-11-3 CAPLUS
CN 2,4(1H,3H)-Pyrimidinedione, 5-fluoro-3-[(4-nitrophenyl)methyl]- (CA INDEX NAME)